

IN THE CLAIMS:

Please amend the claims as follows:

1-134. (Cancelled)

135. (Currently amended): A method of predicting the receptor-modulating activity of a test compound when bound to an estrogen receptor, comprising the steps of:

(1) (a) providing an unliganded estrogen receptor;

(b) contacting thesaid unliganded estrogen receptor with a plurality of reference compounds, ~~said reference compounds~~ known to modulate the biological activity of thesaid estrogen receptor, and wherein each of the unliganded estrogen receptor ~~and the binding of each reference compound to~~ thesaid estrogen receptor bound to each of the reference compounds forms a reference conformation, thesaid plurality of reference compounds selected from the group consisting of estradiol, estriol, nafoxidine, 4-OH tamoxifen, clomifene, premarin, raloxifene, ICI 182,780, 16 α -OH estrone, and progesterone;

(c) contacting the estrogen receptor reference conformations with providing a panel comprising a plurality of peptide conformational probes members representing a plurality of classes, wherein the classes are selected from the group consisting of ER α / β I, ER α / β II, ER α / β III, ER α / β IV, ER α / β V, ER α I, ER α II, ER α III, ER β I, ER β II, and ER β III peptides, wherein thesaid members of thesaid panel possess differential ability to bind to the unliganded reference conformation and one or more of thesaid reference conformations;

(d) ~~contacting said reference conformation with said panel;~~

[[e)] measuring the effect of ~~said reference compound on the binding of~~ thesaid panel members to thesaid estrogen receptor reference conformations, ~~said measuring step forming to form~~ a fingerprint for each member of said plurality of the reference compounds;

(2) (a) providing a test compound;

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(b) contacting thesaid unliganded estrogen receptor with thesaid test compound, wherein the binding of thesaid test compound to thesaid estrogen receptor forms a test conformation;

(c) contacting thesaid test conformation with thesaid panel members;

(d) measuring the ~~effect of said test compound on the binding of~~ thesaid panel members to the test conformation; and

(3) comparing the ~~effect of said test compound on the binding of~~ thesaid panel members to the test conformation and to thesaid reference compound fingerprints to predict the receptor-modulating activity of thesaid test compound when bound to thesaid estrogen receptor.

136.-138 (Canceled)

139. (Currently amended) The method of claim 135, wherein thesaid biological activity of thesaid reference compounds at thesaid estrogen receptor is known for a plurality of different tissues, so that the biological activity of thesaid test compound in thesaid different tissues is predicted.

140-141. (Cancelled).

142. (Currently amended) The method of claim 135, wherein thesaid reference compound is a pharmacological agonist or antagonist of thesaid receptor.

143. (Cancelled)

144. (Currently amended) The method of claim 135, wherein thesaid test compounds are provided ~~and screened~~ in the form of a combinatorial library.

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145. (Currently amended) The method of claim 135, wherein thesaid test compound comprises an organic compound with a molecular weight of less than 500 daltons.

146. (Currently amended) The method of claim 135, wherein thesaid contacting steps are performed in vitro.

147. (Cancelled).

148. (Currently amended) The method of claim 135, wherein at least one of the panel members is a peptide comprising Leu-Xaa-Xaa-Leu-Leu wherein Xaa represents any naturally occurring amino acid.

149. (Currently amended) The method of claim 158[[435]], wherein at least one of the panel members has a substantially higher affinity for the ER α than for the ER β , and at least one other of the panel members has a substantially higher affinity for the ER β than for the ER α .

150. (Currently amended) The method of claim 135, wherein at least one of the panel members binds the estrogen receptor substantially more strongly when the estrogen receptor is bound to estradiol than when the estrogen receptor is not so bound.

151. (Currently amended) The method of claim 135, wherein at least one of the panel members binds the estrogen receptor substantially less strongly when the estrogen receptor is bound to estradiol then when the estrogen receptor it is not so bound.

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152. (Currently amended) The method of claim 158~~[[135]]~~, wherein thesaid panel comprises:

(1) at least one member ~~having with a substantially~~substantially higher affinity for the ER β than for the ER α , and having an ~~whose affinity~~ that is substantially greater for the estradiol-bound ER than for the unliganded ER;

(2) at least one member ~~having with a~~ substantially higher affinity for ER α than for ER β , and having an ~~whose affinity~~ that is substantially the same for the estradiol-bound ER and for the unliganded ER;

(3) at least one member ~~having with a~~ substantially higher affinity for ER α than for ER β , and having an ~~whose affinity~~ that is higher for the estradiol-bound ER α than for the unliganded ER α , and substantially the same for the estradiol-bound ER β and the unliganded ER β ;

(4) at least one member ~~having with a~~ higher affinity for the ER α than for the ER β , and having an ~~whose affinity~~ that is substantially lower for the estradiol bound ER α than for the unliganded ER α , and substantially the same for the estradiol-bound ER β and the unliganded ER β ; and

(5) at least one member ~~having with a~~ substantially higher affinity for the ER β than for the ER α , and having an ~~whose affinity~~ that is substantially lower for the estradiol-bound ER than for the unliganded ER.

153. (Currently amended) The method of claim 135, wherein thesaid reference conformations include ~~a plurality of conformations selected from the group consisting of~~ unliganded receptor, estradiol-liganded receptor, 4-OH tamoxifen liganded receptor, estriol-liganded receptor, nafoxidene-liganded receptor, clomifene-liganded receptor, premarin-liganded receptor, raloxifene-liganded receptor, ICI 182,780-liganded receptor, 16 α -OH estrone-liganded receptor, and progesterone-liganded receptor.

154. (Cancelled).

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155. (Currently amended) The method of claim 135, wherein thesaid method distinguishes among 4-OH tamoxifen, nafoxidene, clomiphene, and raloxifene.

156. (Canceled).

157. (Currently amended) The method of claim 135, wherein at least one member of thesaid panel is a Table 10 peptide, α/β I, α/β II, α/β III, α/β IV, α/β V, α I, α II, α III, β I, β II, and β III, or a peptide[[s]] having the same characterizing binding activity against the reference conformations of the ER as the, ~~and markedly identical to at least one of said~~ Table 10 peptides.

158. (New) The method of claim 135, wherein the estrogen receptor includes an ER α and an ER β .